

# EXHIBIT I1

## Methods

Meta-analysis was performed using the metafor package in R (Version 3.5.1). The rma function was used to apply linear mixed effects models to study results and calculate summary statistics on effect size. Due to varying amounts and types of available data from each included publication, adjusted odds ratios (OR) and standard errors were used as the model inputs. Standard error (SE) was estimated using the relationship: 95% confidence interval = Effect size  $\pm$  1.96\*SE, assuming a roughly normal distribution of data and roughly symmetrical upper and lower confidence interval bounds.

Incorporating adjusted ORs and SE into models in this way provides the added benefit of allowing model use of covariate-adjusted data (versus crude OR data). Weighting was done based on estimates of inverse variance. Study result heterogeneity was estimated based on maximum likelihood methods, and was summarized via an I<sup>2</sup> statistic and associated p-value. The decision to include results from the cohort study by Gertig and colleagues (2000), which reported relative risk (RR), was based on the estimation that the RR value was only nominally different from the OR, a safe assumption in a study sample where less than 0.4% of the cohort developed the condition-of-interest.

## All Papers

## 1. a Any Talc Use and Ovarian CA

**Random Effects Model Results**

Test for Heterogeneity:

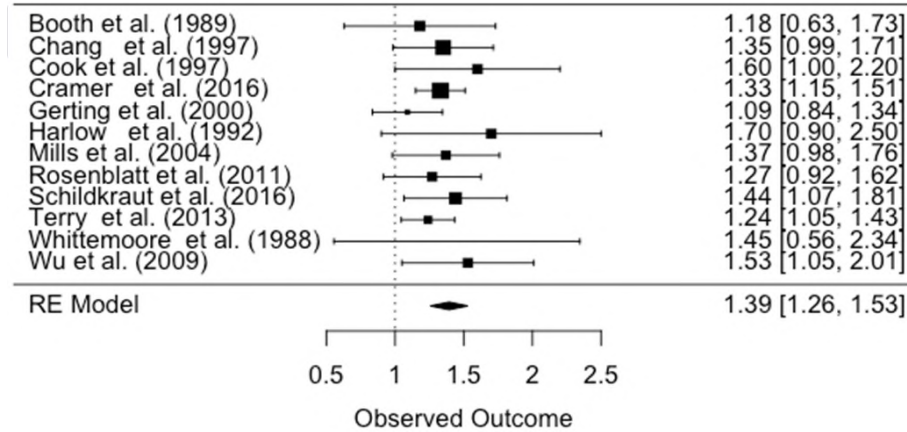
 $I^2$  (total heterogeneity / total variability): 0.0% $Q(df = 10) = 6.8184$ ,  $p\text{-val} = 0.7425$ 

Model Results:

OR estimate 1.3934

Lower Bound 1.2563

Upper bound 1.5305

 $p\text{-value} < .0001$ 

1.39

## 1. b Frequent Talc Use and Ovarian CA

**Random Effects Model Results**

Test for Heterogeneity:

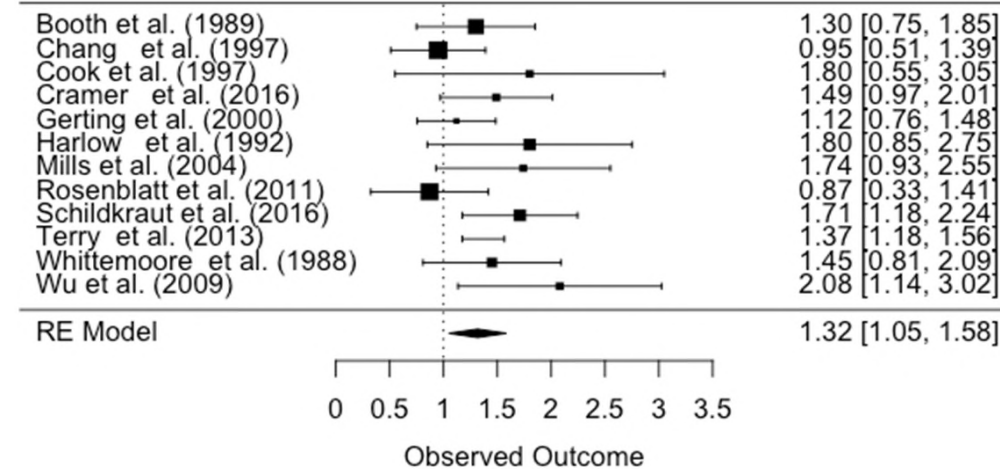
 $I^2$  (total heterogeneity / total variability): 31.87% $Q(df = 10) = 14.1549$ ,  $p\text{-val} = 0.1660$ 

Model Results:

OR estimate 1.316

Lower Bound 1.0502

Upper bound 1.5818

 $p\text{-value} < .0001$ 

1. c Any Talc Use and Serous  
Cancer**Random Effects Model Results**

Test for Heterogeneity:

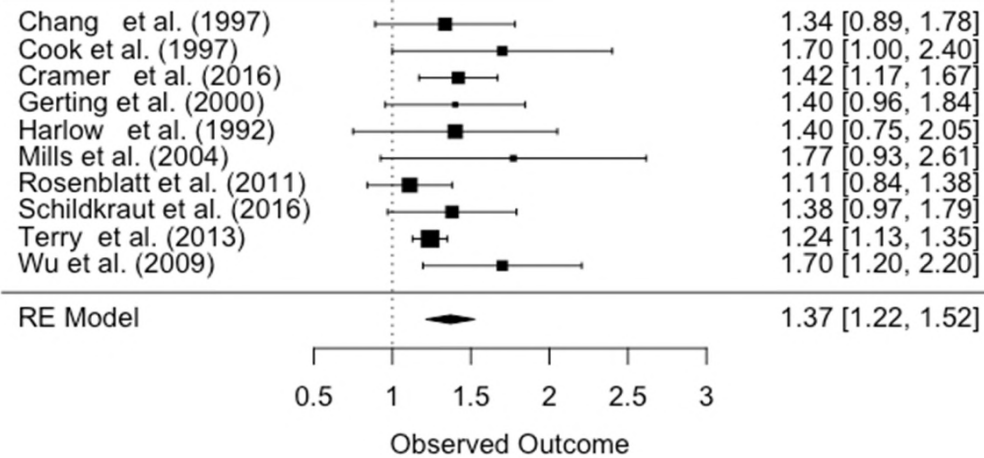
 $I^2$  (total heterogeneity / total variability): 17.08% $Q(df = 9) = 9.1996$ ,  $p\text{-val} = 0.4191$ 

Model Results:

OR estimate 1.3699

Lower Bound 1.216

Upper bound 1.5238

 $p\text{-value} < .0001$ 1. d Frequent Talc use and Serous  
Cancer**Random Effects Model Results**

Test for Heterogeneity:

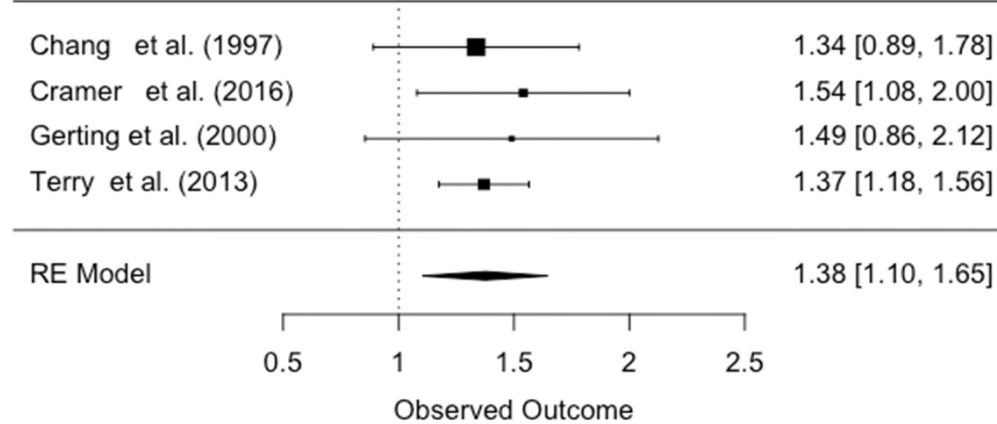
 $I^2$  (total heterogeneity / total variability): 0.00% $Q(df = 3) = 0.5982$ ,  $p\text{-val} = 0.8968$ 

Model Results:

OR estimate 1.3753

Lower Bound 1.1043

Upper bound 1.6463

 $p\text{-value} < .0001$ 

1. e. Frequent Talc use and Serous  
Cancer when available otherwise  
all invasive cancers

#### Random Effects Model Results

Test for Heterogeneity:

$I^2$  (total heterogeneity / total variability): 0.00%

$Q(df = 2) = 0.0167$ ,  $p\text{-val} = 0.9917$

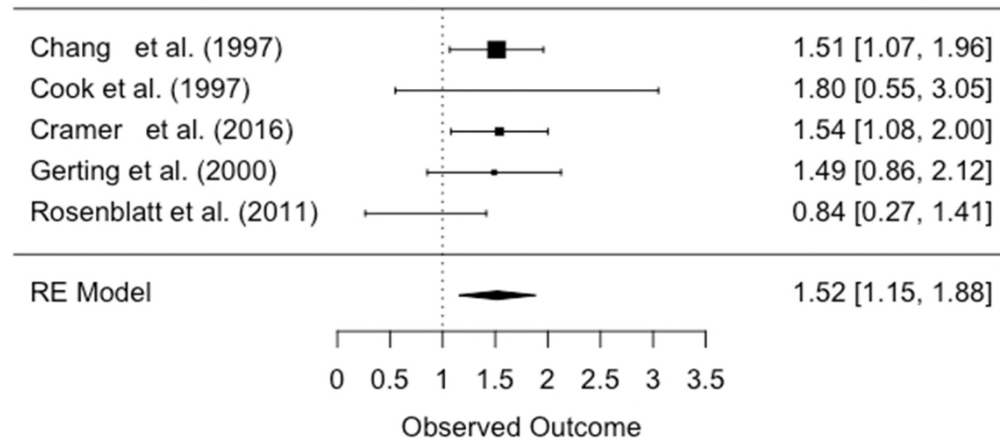
Model Results:

OR estimate 1.5183

Lower Bound 1.152

Upper bound 1.8847

$p\text{-value} < .0001$



Excluding Terry

## 1. a Any Talc Use and Ovarian CA

**Random Effects Model Results**

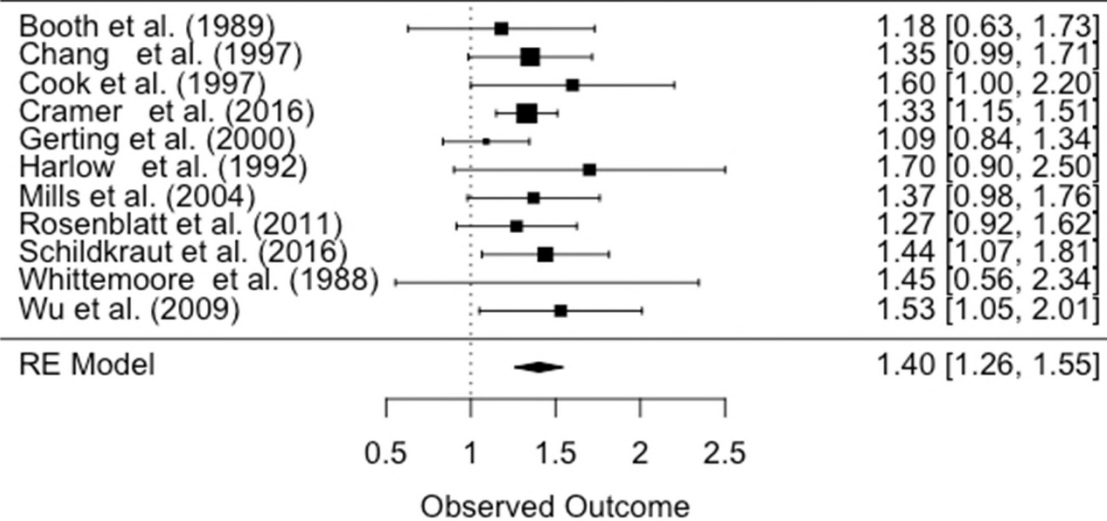
Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 0.00%

Q(df = 9) = 6.3764, p-val = 0.7017

Model Results:

OR estimate 1.402  
 Lower Bound 1.2577  
 Upper bound 1.5463  
 p-value <.0001



## 1. b Frequent Talc Use and Ovarian CA

**Random Effects Model Results**

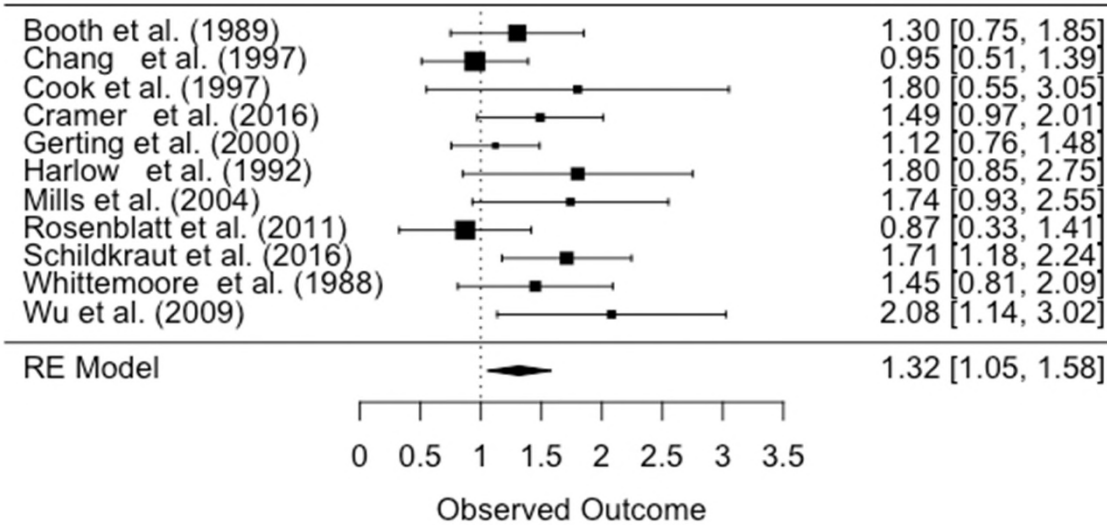
Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 31.87%

Q(df = 10) = 14.1549, p-val = 0.1660

Model Results:

OR estimate 1.316  
 Lower Bound 1.0502  
 Upper bound 1.5818  
 p-value <.0001



ExcludingTerry

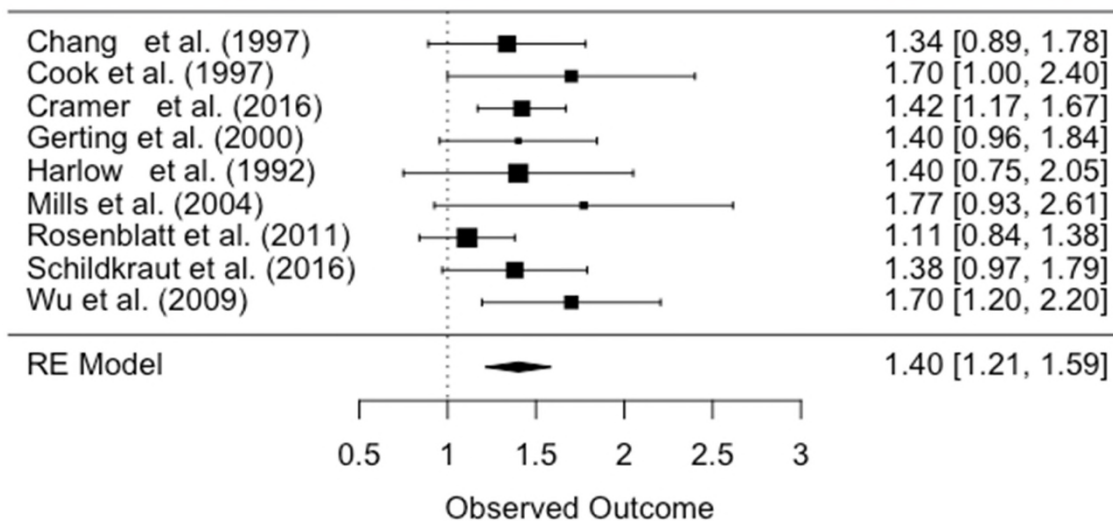
1. c Any Talc Use and Serous  
Cancer**Random Effects Model Results**

Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 12.35%

Q(df = 8) = 7.1202, p-val = 0.5237

OR estimate 1.4009  
 Lower Bound 1.2143  
 Upper bound 1.5874  
 p-value <.0001

1. d Frequent Talc use and Serous  
Cancer**Random Effects Model Results**

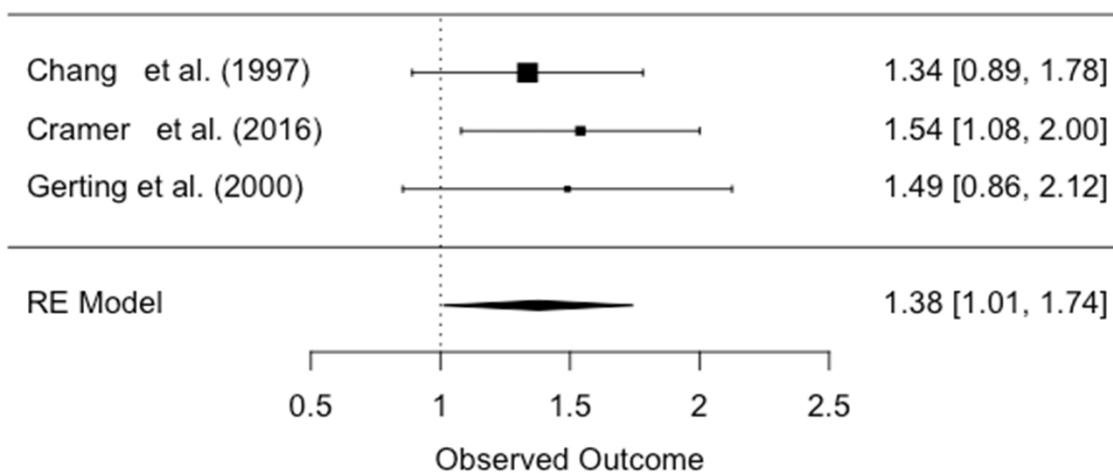
Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 0.00%

Q(df = 2) = 0.4136, p-val = 0.8132

Model Results:

OR estimate 1.3773  
 Lower Bound 1.0109  
 Upper bound 1.7437  
 p-value <.0001



1. e Frequent Talc use and Serous  
Cancer when available otherwise  
all invasive cancers

#### Random Effects Model Results

Test for Heterogeneity:

$I^2$  (total heterogeneity / total variability): 0.00%

$Q(df = 2) = 0.0167$ , p-val = 0.9917

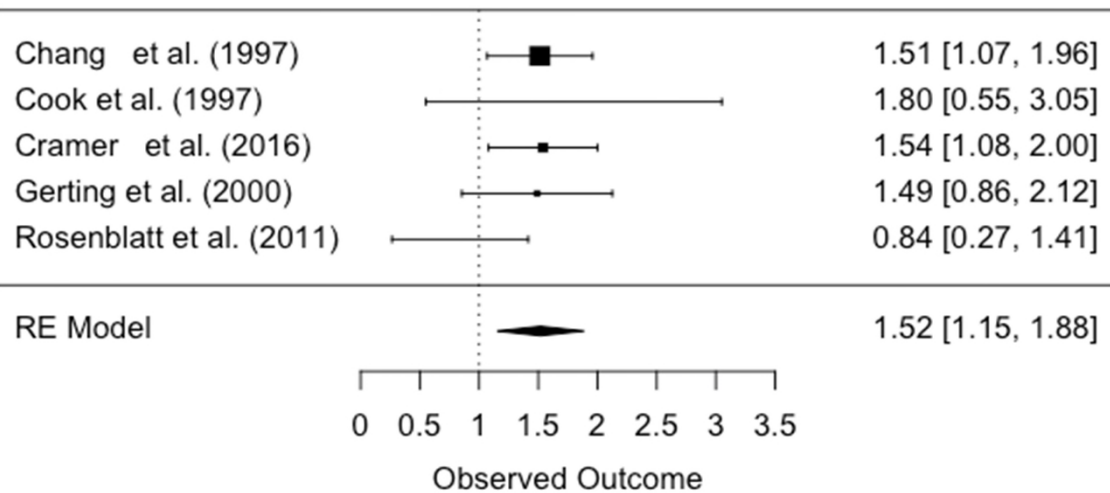
Model Results:

OR estimate 1.5183

Lower Bound 1.152

Upper bound 1.8847

p-value <.0001





Excluding Rosenblatt

## 1. a Any Talc Use and Ovarian CA

**Random Effects Model Results**

Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 0.00%

Q(df = 9) = 6.7928, p-val = 0.6587

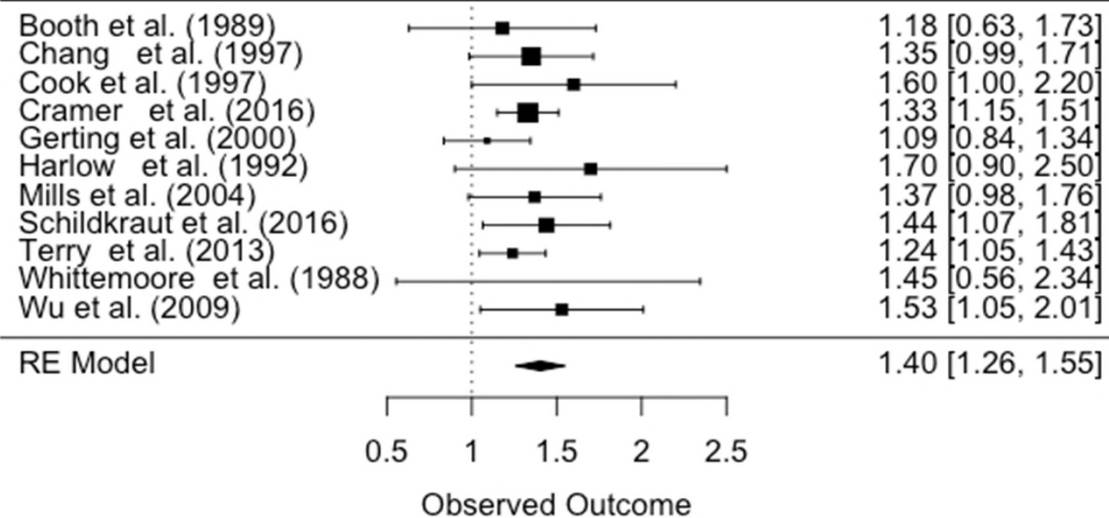
Model Results:

OR estimate 1.4034

Lower Bound 1.258

Upper bound 1.5488

p-value &lt;.0001



## 1. b Frequent Talc Use and Ovarian CA

**Random Effects Model Results**

Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 28.36%

Q(df = 9) = 11.3779, p-val = 0.2507

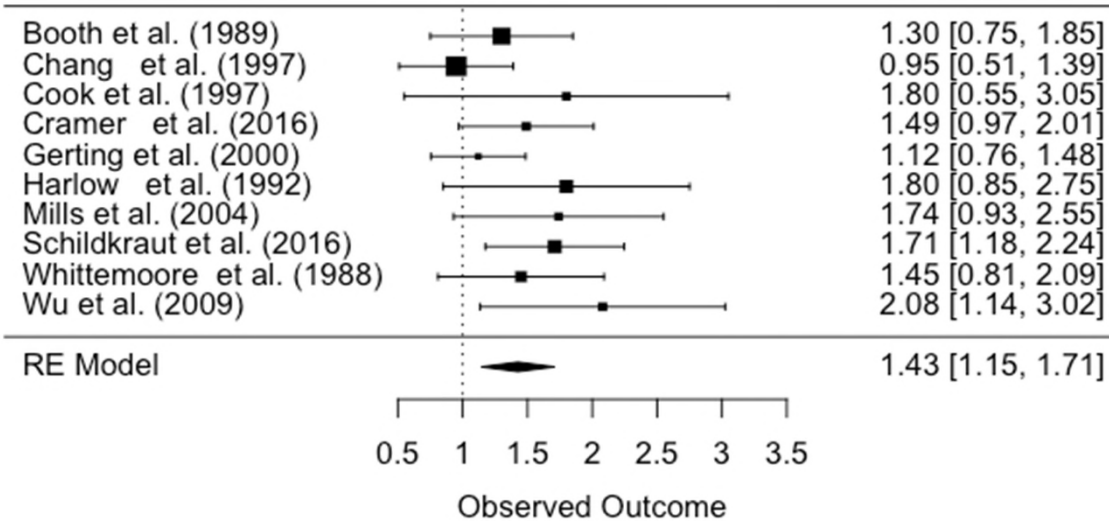
Model Results:

OR estimate 1.4261

Lower Bound 1.1459

Upper bound 1.7062

p-value &lt;.0001



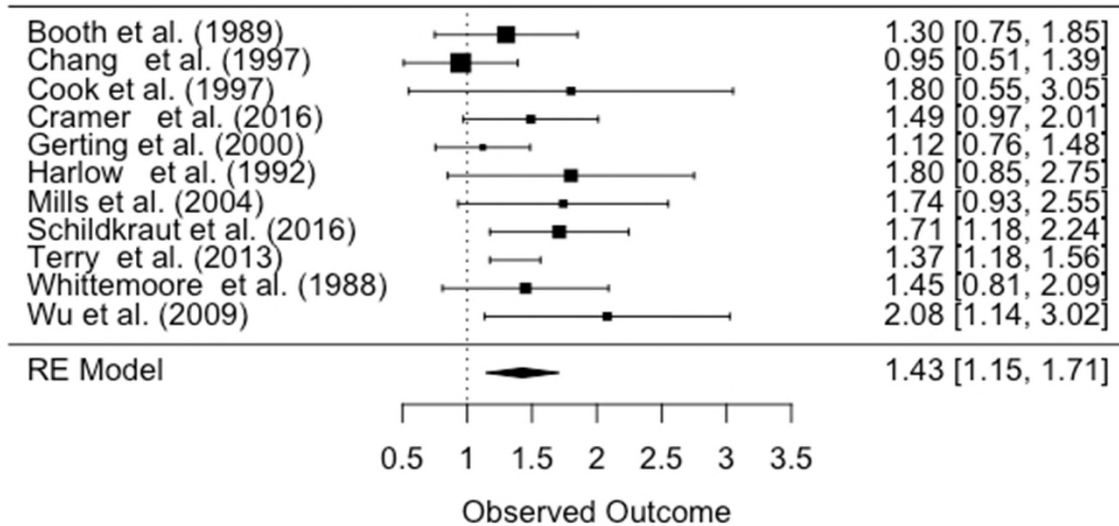
1. c Any Talc Use and Serous  
Cancer**Random Effects Model Results**

Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 28.36% $Q(df = 9) = 11.3779$ ,  $p\text{-val} = 0.2507$ 

Model Results:

OR estimate 1.4261  
 Lower Bound 1.1459  
 Upper bound 1.7062  
 p-value <.0001

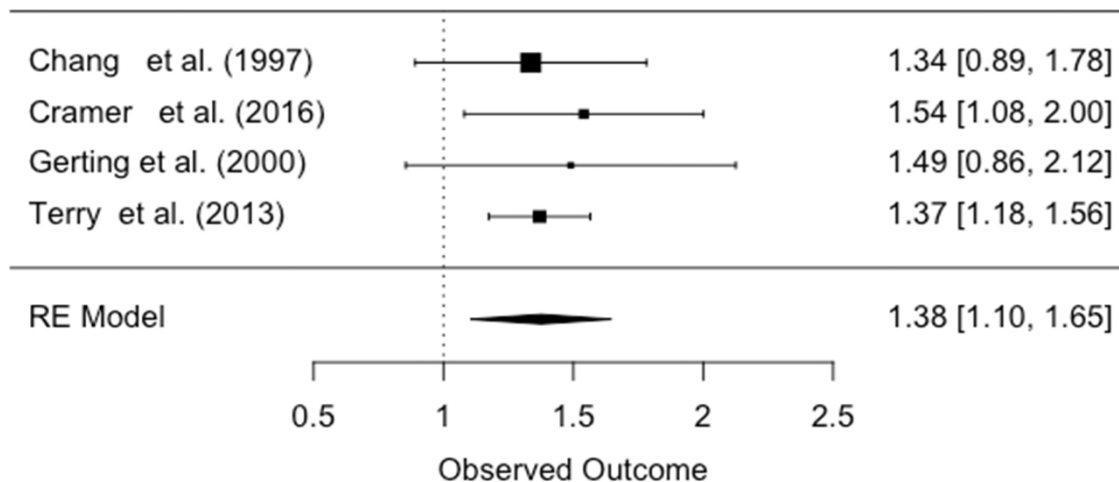
1. d Frequent Talc use and Serous  
Cancer**Random Effects Model Results**

Test for Heterogeneity:

 $I^2$  (total heterogeneity / total variability): 0.00% $Q(df = 3) = 0.5982$ ,  $p\text{-val} = 0.8968$ 

Model Results:

OR estimate 1.3753  
 Lower Bound 1.1043  
 Upper bound 1.6463  
 p-value <.0001



1. e Frequent Talc use and Serous  
Cancer when available otherwise  
all invasive cancers

#### Random Effects Model Results

Test for Heterogeneity:

$I^2$  (total heterogeneity / total variability): 0.00%

$Q(df = 2) = 0.0167$ , p-val = 0.9917

Model Results:

OR estimate 1.5183

Lower Bound 1.152

Upper bound 1.8847

p-value <.0001

